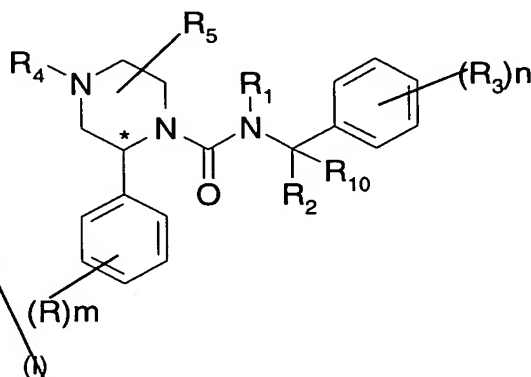


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In the Claims:

Please cancel claims 1 through 19 and in lieu thereof insert claims 20 through 38 as follows.

20. A compound of formula (I)



wherein

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R is a halogen atom or a C₁₋₄ alkyl group;

R₁ is hydrogen or a C₁₋₄ alkyl group;

R₂ is hydrogen, a C₁₋₄ alkyl, C₂₋₆ alkenyl or a C₃₋₇ cycloalkyl group; or R₁ and R₂ together with nitrogen and carbon atom to which they are attached respectively are a 5-6 membered heterocyclic group;

R₃ is a trifluoromethyl, a C₁₋₄ alkyl, a C₁₋₄ alkoxy, a trifluoromethoxy, or a halogen group;

R₄ is hydrogen, a (CH₂)_qR₇ or a (CH₂)_rCO(CH₂)_pR₇ group;

R₅ is hydrogen, a C₁₋₄ alkyl or a COR₆ group;

R₆ is hydrogen, hydroxy, amino, methylamino, dimethylamino, a 5 membered heteroaryl group containing 1 to 3 heteroatoms selected from oxygen, sulphur and nitrogen or a 6 membered heteroaryl group containing 1 to 3 nitrogen atoms;

R₇ is hydrogen, hydroxy or NR₈R₉ wherein R₈ and R₉ are independently hydrogen or C₁₋₄ alkyl optionally substituted by hydroxy, or by amino;

R₁₀ is hydrogen, a C₁₋₄ alkyl group or

R₁₀ together with R₂ is a C₃₋₇ cycloalkyl group;

m is zero or an integer from 1 to 3; n is zero or an integer from 1 to 3; both p and r are independently zero or an integer from 1 to 4; q is an integer from 1 to 4; provided that, when R₁ and R₂ together with nitrogen and carbon atom to which they are attached respectively are a 5 to 6 membered heterocyclic group, i) m is 1 or 2; ii) when m is 1, R is not fluorine and iii) when m is 2, the two substituents R are not both fluorine, and pharmaceutically acceptable salts and solvates thereof.

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21. A compound as claimed in claim 1 wherein n is 2 and R_3 is trifluoromethyl both at the 3 and 5 position.

22. A compound as claimed in claim 1 wherein R is selected independently from halogen or a C_{1-4} alkyl group and m is 1 or 2.

23. A compound as claimed in claim 1 wherein m is 2, R is selected independently from halogen or methyl group at 2 or 4 position.

24. A compound as claimed in claim 1 wherein R_5 is hydrogen or a methyl group.

25. A compound as claimed in claim 1 wherein R_1 is hydrogen or a methyl group.

26. A compound as claimed in claim 1 wherein R_4 is hydrogen, a $(CH_2)_rCO(CH_2)_pR_7$ or $CH_2)_qR_7$ group, wherein R_7 represents an amine, both p and r are independently zero or 1; and q is 1 or 2.

27. A compound of formula (I) as claimed in claim 1 wherein R is selected independently from halogen or methyl, R_3 is trifluoromethyl both at the 3 and 5 position, R_1 is hydrogen or methyl, R_2 is hydrogen, methyl, 2-propenyl or a cyclopropyl group or together with R_1 is a 3,6-dihydro-2H-pyridin-1-yl, a piperidin-1-yl or a pyrrolidin-1-yl group, R_{10} represents hydrogen, a methyl or R_{10} together with R_2 is a cyclopropyl group, R_4 is hydrogen, an aminoacetyl or amino ethyl group and R_5 is hydrogen or a methyl group.

28. A compound of formula (I) as claimed in claim 1 wherein R is selected independently from halogen or methyl and m is 2, R_3 is trifluoromethyl both at the 3 and 5 position, R_1 and R_2 are independently hydrogen or methyl, R_4 is hydrogen and R_5 is hydrogen.

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29. A compound selected from:

2-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide;

2-(2-isopropyl-phenyl)-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide;

2-(4-fluoro-3-methyl-phenyl)-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide;

2-(2,4-difluoro-phenyl)-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide;

2-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid [1-(3,5-bis-trifluoromethyl-phenyl)ethyl]-methyl-amide;

2-(4-fluoro-phenyl)-piperazine-1-carboxylic acid (3,4-bis-trifluoromethyl-benzyl)-methyl-amide;

2-phenyl-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide;

2-(2,4-dichloro-phenyl)-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide;

2-(3,4-dichloro-phenyl)-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide;

2-(4-fluoro-2-methyl-phenyl)-3-methyl-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide;

2-(2-methyl-4-fluoro-phenyl)-6-methyl-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide;

2-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid [1-(3,5-bis-trifluoromethyl-phenyl)ethyl]-methyl-amide;

4-(2-amino-acetyl)-2-(S)-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide ;

2-(S)-(4-fluoro-2-methyl-phenyl)-4-(piperidine-4-carbonyl)-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide;

4-(2-amino-ethyl)-2-(S)-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide;

2-(S)-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid [(1-3,5-bis-trifluoromethyl-phenyl)-cyclopropyl]-methyl-amide;

[2-(3,5-bis-trifluoromethyl-phenyl)-pyrrolidin-1-yl]-[2-(S)-(4-fluoro-2-methyl-phenyl)-piperazin-1-yl]-methanone;

[2-(3,5-bis-trifluoromethyl-phenyl)-3,6-dihydro-2H-pyridin-1-yl]-[2-(S)-(4-fluoro-2-methyl-phenyl)-piperazin-1-yl]-methanone;

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2-(3,5-bis-trifluoromethyl-phenyl)-piperidin-1-yl]-[2-(S)-(4-fluoro-2-methyl-phenyl)-piperazin-1-yl]-methanone;

2-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid [1-(3,5-bis-trifluoromethyl-phenyl)-but-3-enyl]-methyl-amide;

2-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid [1-(3,5-bis-trifluoromethyl-phenyl)-2-methyl-propyl]-methyl-amide;

2-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid [(3,5-bis-trifluoromethyl-phenyl)-cyclopropyl-methyl]-methyl-amide;

and enantiomers, pharmaceutically acceptable salts, and solvates thereof.

30. 2-(S)-(4-fluoro-2-methyl-phenyl)-4-(piperidine-4-carbonyl)-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide hydrochloride.

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31. 4-(2-amino-acetyl)-2-(S)-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid (3,5-bis-trifluoromethyl-benzyl)-methyl-amide hydrochloride.

32. 2-(S)-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid ~~1-(R)-(3,5-bis-trifluoromethyl-phenyl)-ethyl]-methyl-amide ethanesulphonate.~~

33. 2-(S)-(4-fluoro-2-methyl-phenyl)-piperazine-1-carboxylic acid [1-(R)-(3,5-bis-trifluoromethyl-phenyl)-ethyl]-methyl-amide acetate.

34. A pharmaceutical composition comprising a compound as claimed in claim 1 in admixture with one or more physiologically acceptable carriers or excipients.

~~35. A method for the treatment of a mammal, in particular in the treatment of conditions mediated by tachykinins comprising administration of an effective amount of a compound claimed in of any claims 20 to 33.~~

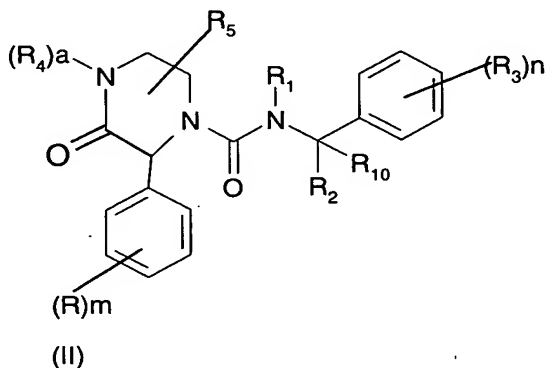
36. The method of Claim 35 wherein said tachykinins is substance P or other neurokinins.

37. The method of Claim 35 wherein said mammal is man.

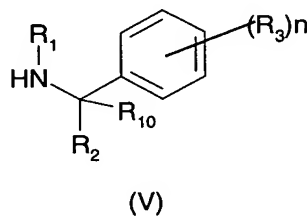
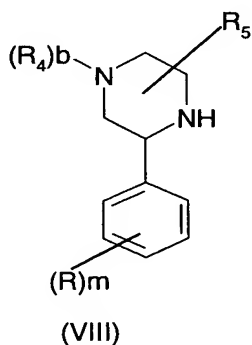
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38. A process (A) for the preparation of a compound of formula (I) as claimed in claim 1, wherein R_4 is hydrogen or a $(CH_2)_q R_7$ group, provided that when R_5 is a C_{1-4} alkyl or a COR_6 group, R_5 is not in the 3 position of the piperazine ring, which comprises reduction of a compound of formula (II), wherein $(R_4)_a$ is hydrogen or a suitable nitrogen protecting group or $(R_4)_a$ is a $(CH_2)_q R_7$ group or protecting derivatives thereof; or



a process (B) for the preparation of a compound of formula (I) as claimed in claim 1, wherein R_4 is hydrogen or a $(CH_2)_r CO(CH_2)_p R_7$ group which comprises the reaction of a compound of formula (VIII), wherein $(R_4)_b$ represents a nitrogen protecting group or $(R_4)_b$ is $(CH_2)_r CO(CH_2)_p R_7$ or a protecting group thereof with triphosgene and an organic base followed by addition of the amine (V)



followed where necessary or desired by one or more of the following steps:

- (i) removal of any protecting group;
- (ii) isolation of the compound as salt thereof;
- (iii) separation of a compound of formula (I) or derivative thereof into the enantiomers thereof.--